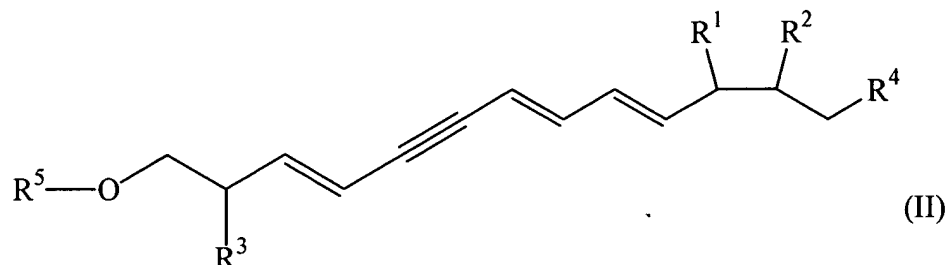


What is claimed is:

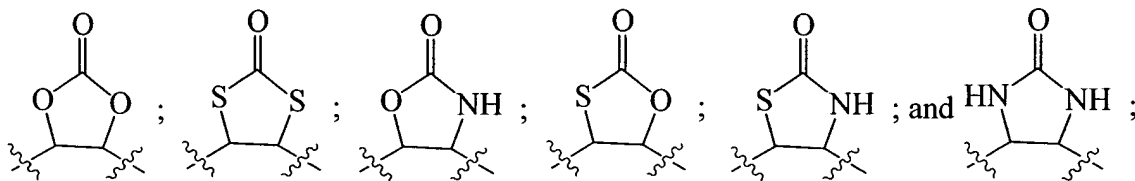
1. A compound of formula (II):



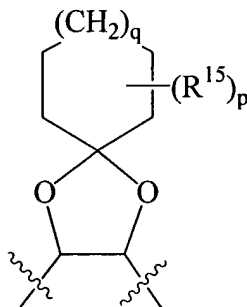
wherein:

each R^1 , R^2 and R^3 are independently halo, $-OR^6$, $-SR^6$, $-S(O)_tR^7$ (where t is 1 or 2) or $-N(R^7)R^8$; or R^1 and R^2 together with the carbons to which they are attached form a monocyclic

heterocyclic structure selected from the following:



or R^1 and R^2 together with the carbons to which they are attached form the following bicyclic heterocyclic structure:



(where q is 0 to 3, p is 1 to 4 and each R^{15} is hydrogen, alkyl, aralkyl or aryl);

each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;

each R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl,

alkoxy, halo, haloalkyl and haloalkoxy);

each R^6 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(S)R^7$, $-C(O)OR^{14}$, $-C(S)OR^{14}$, $-C(O)N(R^7)R^8$, or $-C(S)N(R^7)R^8$;

each R^7 is independently hydrogen, alkyl, cycloalkyl, aryl, or aralkyl;

R^8 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(O)OR^{14}$, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);

each R^9 is independently a direct bond or a straight or branched alkylene chain;

each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;

each R^{11} is independently $-C(O)OR^7$, $-C(O)N(R^7)_2$, $-P(O)(OR^7)_2$, $-S(O)_2OR^7$, $-S(O)_2N(H)R^7$ or tetrazole;

R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);

R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene; and

R^{14} is alkyl, aryl or aralkyl;

as a single stereoisomer, a mixture of stereoisomers, a racemic mixture of stereoisomers; or as a cyclodextrin clathrate thereof, or as a pharmaceutically acceptable salt thereof.

2. The compound of Claim 1 wherein:

R^1 , R^2 and R^3 are each independently halo, $-OR^6$, $-SR^6$ or $-N(R^7)R^8$;

each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;

R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, and haloalkoxy);

each R^6 is independently hydrogen, alkyl, aralkyl, $-C(O)R^7$ or $-C(O)OR^7$;

each R^7 is independently hydrogen, alkyl, aryl, or aralkyl;

R^8 is independently hydrogen, alkyl, aryl, aralkyl, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);

each R^9 is independently a direct bond or a straight or branched alkylene chain;

each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;

each R^{11} is independently $-C(O)OR^7$ or $-C(O)N(R^7)_2$;

R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo and haloalkoxy);

R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene.

3. The compound of Claim 2 wherein:

R^1 , R^2 and R^3 are each independently halo, $-OR^6$, or $-SR^6$;

R^4 is $-R^9-O-R^{10}-R^{11}$;

R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, and haloalkoxy);

each R^6 is independently hydrogen, alkyl, aryl, or aralkyl;

each R^7 is independently hydrogen, alkyl, aryl, or aralkyl;

R^9 is a direct bond or a straight or branched alkylene chain;

R^{10} is an straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene; and

R^{11} is $-C(O)OR^7$ or $-C(O)N(R^7)_2$.

4. The compound of Claim 3 wherein:

R^1 , R^2 and R^3 are each $-OR^6$;

R^4 is $-R^9-O-R^{10}-R^{11}$;

R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting

of alkyl, alkoxy, halo, and haloalkoxy);

R^6 is hydrogen, alkyl, aryl, or aralkyl;

each R^7 is independently hydrogen, alkyl, aryl, or aralkyl;

R^9 is a direct bond;

R^{10} is a straight or branched alkylene chain, a straight or branched alkenylene chain, or a straight or branched alkynylene chain; and

R^{11} is $-C(O)OR^7$ or $-C(O)N(R^7)_2$.

5. The compound of Claim 4 selected from the group consisting of the following:

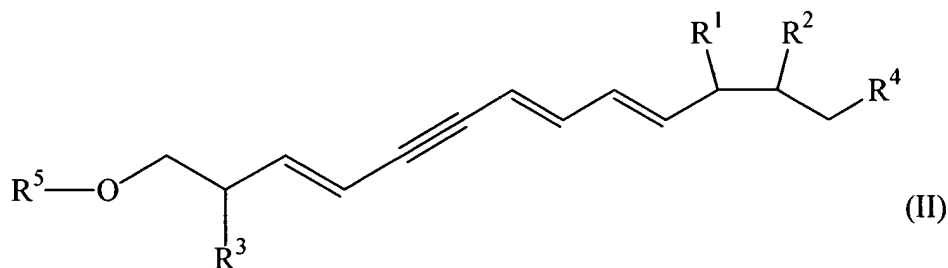
(5*S*,6*R*,7*E*,9*E*,13*E*,15*S*)-16-(4-fluorophenoxy)-5,6,15-trihydroxy-3-oxahexadeca-7,9,13-trien-11-ynoic acid, methyl ester;

(5*S*,6*R*,7*E*,9*E*,13*E*,15*S*)-16-(4-fluorophenoxy)-5,6,15-trihydroxy-3-oxahexadeca-7,9,13-trien-11-ynoic acid;

(5*S*,6*S*,7*E*,9*E*,13*E*,15*S*)-16-(4-fluorophenoxy)-5,6,15-trihydroxy-3-oxahexadeca-7,9,13-trien-11-ynoic acid, methyl ester; and

(5*S*,6*S*,7*E*,9*E*,13*E*,15*S*)-16-(4-fluorophenoxy)-5,6,15-trihydroxy-3-oxahexadeca-7,9,13-trien-11-ynoic acid.

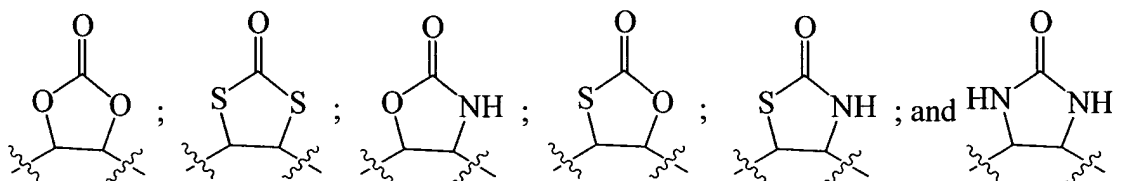
6. A pharmaceutical composition useful in treating an inflammatory or autoimmune disorder in a mammal, which composition comprises one or more pharmaceutically acceptable excipient(s) and a therapeutically effective amount of a compound of formula (II):



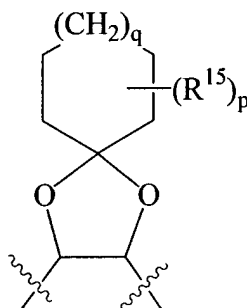
wherein:

each R^1 , R^2 and R^3 are independently halo, $-OR^6$, $-SR^6$, $-S(O)_tR^7$ (where t is 1 or 2) or $-N(R^7)R^8$; or R^1 and R^2 together with the carbons to which they are attached form a monocyclic

heterocyclic structure selected from the following:



or R^1 and R^2 together with the carbons to which they are attached form the following bicyclic heterocyclic structure:



- (where q is 0 to 3, p is 1 to 4 and each R^{15} is hydrogen, alkyl, aralkyl or aryl);
- each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;
- each R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);
- each R^6 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(S)R^7$, $-C(O)OR^{14}$, $-C(S)OR^{14}$, $-C(O)N(R^7)R^8$, or $-C(S)N(R^7)R^8$;
- each R^7 is independently hydrogen, alkyl, cycloalkyl, aryl, or aralkyl;
- R^8 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(O)OR^{14}$, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);
- each R^9 is independently a direct bond or a straight or branched alkylene chain;
- each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;
- each R^{11} is independently $-C(O)OR^7$, $-C(O)N(R^7)_2$, $-P(O)(OR^7)_2$, $-S(O)_2OR^7$, $-S(O)_2N(H)R^7$ or

tetrazole;

R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);

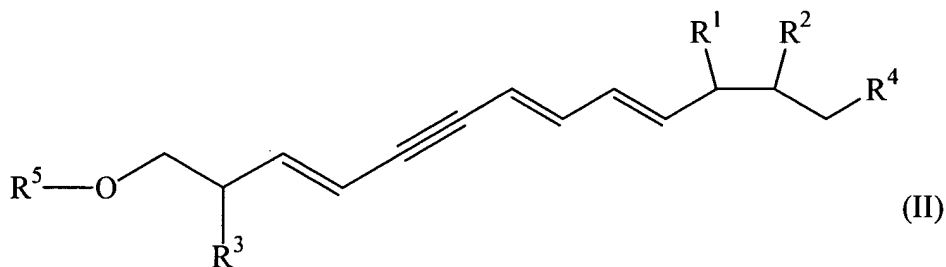
R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene; and

R^{14} is alkyl, aryl or aralkyl;

as a single stereoisomer, a mixture of stereoisomers, a racemic mixture of stereoisomers; or as a cyclodextrin clathrate thereof, or as a pharmaceutically acceptable salt thereof.

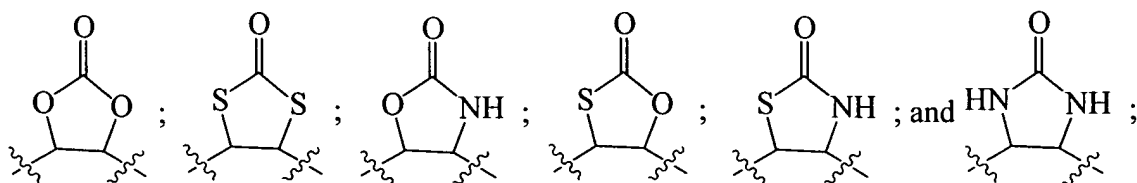
7. The pharmaceutical composition of Claim 6 wherein the mammal is a human.

8. A pharmaceutical composition useful in treating pulmonary or respiratory tract inflammation in a mammal, wherein the composition comprises one or more pharmaceutically acceptable excipient(s) and a therapeutically effective amount of a compound of formula (II):

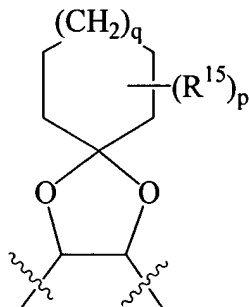


wherein:

each R^1 , R^2 and R^3 are independently halo, $-OR^6$, $-SR^6$, $-S(O)_tR^7$ (where t is 1 or 2) or $-N(R^7)R^8$; or R^1 and R^2 together with the carbons to which they are attached form a monocyclic heterocyclic structure selected from the following:



or R^1 and R^2 together with the carbons to which they are attached form the following bicyclic heterocyclic structure:



- (where q is 0 to 3, p is 1 to 4 and each R^{15} is hydrogen, alkyl, aralkyl or aryl);
- each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;
- each R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);
- each R^6 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(S)R^7$, $-C(O)OR^{14}$, $-C(S)OR^{14}$, $-C(O)N(R^7)R^8$, or $-C(S)N(R^7)R^8$;
- each R^7 is independently hydrogen, alkyl, cycloalkyl, aryl, or aralkyl;
- R^8 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(O)OR^{14}$, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);
- each R^9 is independently a direct bond or a straight or branched alkylene chain;
- each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;
- each R^{11} is independently $-C(O)OR^7$, $-C(O)N(R^7)_2$, $-P(O)(OR^7)_2$, $-S(O)_2OR^7$, $-S(O)_2N(H)R^7$ or tetrazole;

R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);

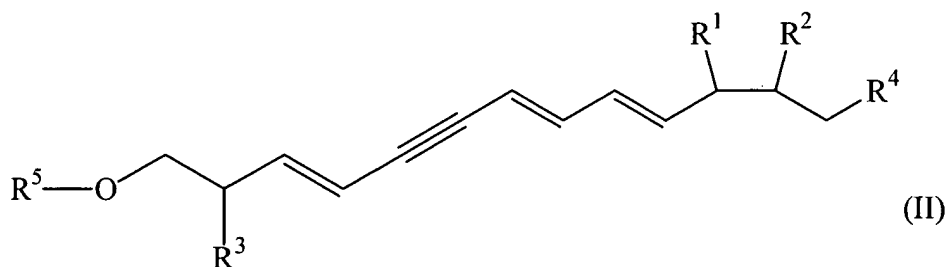
R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene; and

R^{14} is alkyl, aryl or aralkyl;

as a single stereoisomer, a mixture of stereoisomers, a racemic mixture of stereoisomers; or as a cyclodextrin clathrate thereof, or as a pharmaceutically acceptable salt thereof.

9. The pharmaceutical composition of Claim 8 wherein the mammal is a human.

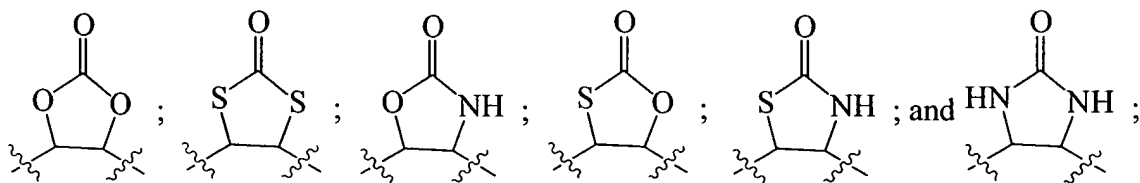
10. A method of treating an inflammatory or autoimmune disorder in a mammal, wherein the method comprises administering to the mammal in need thereof a therapeutically effective amount of a compound of formula (II):



wherein:

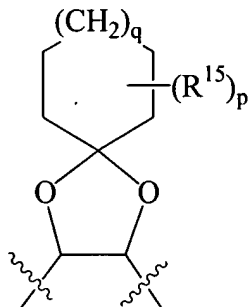
each R^1 , R^2 and R^3 are independently halo, $-OR^6$, $-SR^6$, $-S(O)_tR^7$ (where t is 1 or 2) or $-N(R^7)R^8$; or R^1 and R^2 together with the carbons to which they are attached form a monocyclic

heterocyclic structure selected from the following:



or R^1 and R^2 together with the carbons to which they are attached form the following bicyclic

heterocyclic structure:



- (where q is 0 to 3, p is 1 to 4 and each R^{15} is hydrogen, alkyl, aralkyl or aryl);
- each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;
- each R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);
- each R^6 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(S)R^7$, $-C(O)OR^{14}$, $-C(S)OR^{14}$, $-C(O)N(R^7)R^8$, or $-C(S)N(R^7)R^8$;
- each R^7 is independently hydrogen, alkyl, cycloalkyl, aryl, or aralkyl;
- R^8 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(O)OR^{14}$, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);
- each R^9 is independently a direct bond or a straight or branched alkylene chain;
- each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;
- each R^{11} is independently $-C(O)OR^7$, $-C(O)N(R^7)_2$, $-P(O)(OR^7)_2$, $-S(O)_2OR^7$, $-S(O)_2N(H)R^7$ or tetrazole;
- R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);

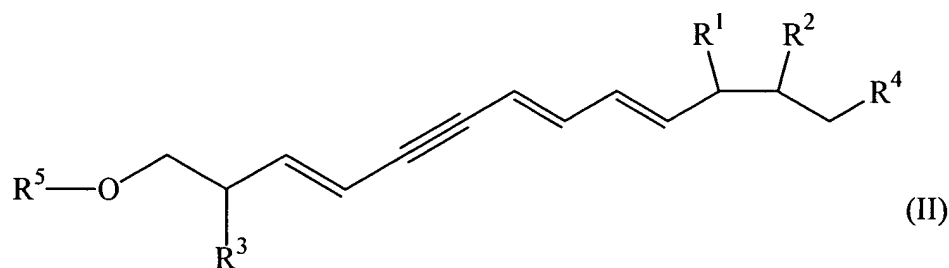
R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene; and R^{14} is alkyl, aryl or aralkyl;

as a single stereoisomer, a mixture of stereoisomers, a racemic mixture of stereoisomers; or as a cyclodextrin clathrate thereof, or as a pharmaceutically acceptable salt thereof.

11. The method of Claim 10 wherein the mammal is a human.

12. The method of Claim 11 wherein the inflammatory or autoimmune disorder is selected from the group consisting of the following:
allergic contact dermatitis, allergic rhinitis, chemical and non-specific irritant contact dermatitis, urticaria, atopic dermatitis, psoriasis, acute myocardial ischemia and infarction, acute hemorrhagic or ischemic stroke, multiple sclerosis, rheumatoid arthritis, osteoarthritis and systemic lupus erythematosus, acute and chronic organ transplant rejection, transplant arteriosclerosis and fibrosis, hypertension, atherosclerosis, aneurysm, critical leg ischemia, peripheral arterial occlusive disease, Reynaud's syndrome, diabetic nephropathy, diabetic neuropathy, and diabetic retinopathy, delayed neurodegeneration in stroke, Alzheimer's disease, Parkinson's disease, benign prostatic hyperplasia, leukemia, lymphoma, prostate cancer, breast cancer, lung cancer, malignant melanoma, renal carcinoma, head and neck tumors and colorectal cancer.

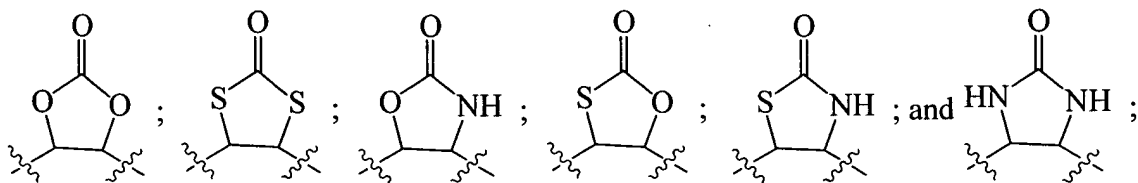
13. A method of treating pulmonary or respiratory tract inflammation in a mammal, wherein the method comprises administering to the mammal in need thereof a therapeutically effective amount of a compound of formula (II):



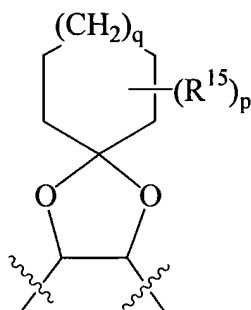
wherein:

each R^1 , R^2 and R^3 are independently halo, $-OR^6$, $-SR^6$, $-S(O)_tR^7$ (where t is 1 or 2) or $-N(R^7)R^8$;

or R^1 and R^2 together with the carbons to which they are attached form a monocyclic heterocyclic structure selected from the following:



or R^1 and R^2 together with the carbons to which they are attached form the following bicyclic heterocyclic structure:



- (where q is 0 to 3, p is 1 to 4 and each R^{15} is hydrogen, alkyl, aralkyl or aryl);
- each R^4 is $-R^9-R^{12}$, $-R^9-R^{13}-R^{11}$, $-R^9-O-R^{10}-R^{11}$, $-R^9-O-R^{12}$, $-R^9-C(O)-R^{10}-R^{11}$, $-R^9-N(R^7)-R^{10}-R^{11}$, $-R^9-S(O)_t-R^{10}-R^{11}$ (where t is 0 to 2), or $-R^9-C(F)_2-R^9-R^{11}$;
- each R^5 is aryl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (optionally substituted by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);
- each R^6 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(S)R^7$, $-C(O)OR^{14}$, $-C(S)OR^{14}$, $-C(O)N(R^7)R^8$, or $-C(S)N(R^7)R^8$;
- each R^7 is independently hydrogen, alkyl, cycloalkyl, aryl, or aralkyl;
- R^8 is independently hydrogen, alkyl, aryl, aralkyl, $-C(O)R^7$, $-C(O)OR^{14}$, or cycloalkyl (optionally substituted with one more substituents selected from the group consisting of alkyl, $-N(R^7)_2$, and $-C(O)OR^7$);
- each R^9 is independently a direct bond or a straight or branched alkylene chain;
- each R^{10} is independently a straight or branched alkylene chain, a straight or branched alkenylene chain, a straight or branched alkynylene chain or a cycloalkylene;

each R^{11} is independently $-C(O)OR^7$, $-C(O)N(R^7)_2$, $-P(O)(OR^7)_2$, $-S(O)_2OR^7$, $-S(O)_2N(H)R^7$ or tetrazole;

R^{12} is aryl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy) or aralkyl (substituted by $-C(O)OR^7$ or $-C(O)N(R^7)_2$ and optionally by one or more substituents selected from the group consisting of alkyl, alkoxy, halo, haloalkyl and haloalkoxy);

R^{13} is a branched alkylene chain, a straight or branched alkenylene chain or a cycloalkylene; and

R^{14} is alkyl, aryl or aralkyl;

as a single stereoisomer, a mixture of stereoisomers, a racemic mixture of stereoisomers; or as a cyclodextrin clathrate thereof, or as a pharmaceutically acceptable salt thereof.

14. The method of Claim 13 wherein the mammal is a human.

15. The method of Claim 10 wherein the inflammatory or autoimmune disorder is selected from the group consisting of the following:

septic or endotoxic shock, hemorrhagic shock, shock-like syndromes, capillary leak syndrome induced by cancer immunotherapy, acute respiratory distress syndrome, traumatic shock, immune- and pathogen-induced pneumonias, immune-complex-mediated pulmonary injury, immune-complex-mediated chronic obstructive pulmonary disease, inflammatory bowel disease, acute renal failure, ischemic bowel disease, immune-complex-mediated glomerulonephritis, insulin-dependent diabetes mellitus, ocular disorders, HIV dementia, encephalitis, inflammatory and neuropathic pain, periodontal disease, and ear infections.

16. The method of Claim 15 wherein the inflammatory or autoimmune disorder is an inflammatory bowel disease selected from the group consisting of Crohn's disease, ulcerative colitis and gastrointestinal ulcers.

17. The method of Claim 16 wherein the inflammatory or autoimmune disorder is Crohn's disease.